# Package 'lrgpr'

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|--|
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| <b>Description</b> Fit a Low Rank Gaussian Process Regression (LRGPR) / Linear Mixed Model (LMM) for large datasets. These models are widely used in statistical genetics as a test of association while correcting for the confounding effects of kinship and population structure. |
| <b>Depends</b> R (>= 3.0.0), methods, Rcpp, RcppGSL, RcppProgress, MASS,parallel, doParallel, formula.tools, BH, bigmemory (>= 4.4.7),biganalytics, and  |
| LinkingTo Rcpp, RcppGSL  |
| <pre>URL http://lrgpr.r-forge.r-project.org/</pre>   |
| License GPL (>= 2)   |
| Collate 'genericFunctions.R' 'lrgpr.R' 'plink.R' 'plots.R'   |
| R topics documented:   |
| BIC.lrgpr coefficients.lrgpr convertToBinary cooks.distance.lrgpr criterion.lrgpr cv.lrgpr df.residual.lrgpr error.bar getAlleleFreq getAlleleVariance   |

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AIC.lrgpr

Akaike's Information Criterion (AIC)

# Description

AIC for model fit by lrgpr

# Usage

```
AIC.lrgpr(object, ..., k = 2)
```

# Arguments

```
object model fit with lrgpr
... other arguments
```

k for compotability, not used

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BIC.lrgpr

Bayesian Information Criterion (BIC)

# Description

```
BIC for model fit by {\tt lrgpr}
```

# Usage

```
BIC.lrgpr(object, ...)
```

# Arguments

```
object model fit with lrgpr
... other arguments
```

```
coefficients.lrgpr Extract Model Coefficients
```

# Description

Coefficients estimated with lrgpr

# Usage

```
coefficients.lrgpr(object)
```

```
object model fit with lrgpr
... other arguments
```

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convertToBinary

Convert ASCII to binary file

#### Description

'convertToBinary' converts TPED/DOSAGE/GEN files to binary format

#### Usage

```
convertToBinary(filename, filenameOut, format,
  nthreads = detectCores(logical = TRUE))
```

#### **Arguments**

```
filename file to be converted

filenameOut name of binary file produced

format specify 'TPED', 'DOSAGE' or 'GEN'
```

#### **Details**

- TPED: plink file can be in either -recode or -recode12 format
- DOSAGE: file follows plink format: http://pngu.mgh.harvard.edu/~purcell/plink/dosage.shtml

#### Example:

```
SNP A1 A2 F1 I1 F2 I2 F3 I3
rs0001 A C 0.98 0.02 1.00 0.00 0.00 0.01
rs0002 G A 0.00 1.00 0.00 0.00 0.99 0.01
where the F* values correspond to the dosage values
```

• GEN: file follow OXFORD format

```
cooks.distance.lrgpr
```

Regression Deletion Diagnostics

### Description

Basic quantities for regression deletion diagnostics from fit of lrgpr

#### Usage

```
cooks.distance.lrgpr(model,
  infl = lm.influence(model, do.coef = FALSE),
  res = weighted.residuals(model),
  sd = sqrt(deviance(model)/df.residual(model)),
  hat = infl$hat, ...)
```

criterion.lrgpr 5

#### **Arguments**

| model | model fit with lrgpr                            |
|-------|---|
| infl  | influence structure as returned by lm.influence |
| res   | residuals                                       |
| sd    | standard deviation to use                       |
| hat   | hat values                                      |
|       | other arguments                                 |
|       |   |

criterion.lrgpr

Compute AIC/BIC/GCV for lrgpr() model as rank changes

### **Description**

'criterion.lrgpr' evaluate information criteria to select an optimal rank

# Usage

```
criterion.lrgpr(formula, features, order, rank = c(seq(1, 10), seq(20, 100, by = 10)), seq(200, 1000, by = 100)))
```

## Arguments

formula standard linear modeling syntax as used in 'lm'

features matrix from which the SVD is performed

order sorted indices of features. When rank is 10, decomp = svd(X[,order[1:10]])

rank array with elements indicating the number of confounding covariates to be used in the random effect.

#### See Also

```
plot.criterion.lrgpr, cv.lrgpr
#'
```

#### **Examples**

```
n = 300
p = 5000
X = matrix(sample(0:2, n*p, replace=TRUE), nrow=n)
dcmp = svd(X)
# simulate response
h_sq = .8
eta = dcmp$u[,1:2] %*% rgamma(2, 2, 1)
error_var = (1-h_sq) / h_sq * var(eta)
```

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```
y = eta + rnorm(n, sd=sqrt(error_var))
# Get ordering based on marginal correlation
i = order(cor(y, X)^2, decreasing=TRUE)
# Fit AIC / BIC / GCV based on degrees of freedom
fit = criterion.lrgpr( y ~ 1, features=X, order=i)
plot.criterion.lrgpr(fit)
```

cv.lrgpr

Cross-validation for LRGPR

#### **Description**

'cv.lrgpr' fits cross-validation for multiple ranks of the LRGPR

# Usage

```
cv.lrgpr(formula, features, order, nfolds = 10,

rank = c(seq(0, 10), seq(20, 100, by = 10), seq(200, 1000, by = 100)),

nthreads = 1)
```

#### **Arguments**

formula standard linear modeling syntax as used in 'lm'

features matrix from which the SVD is performed

order sorted indices of features. When rank is 10, decomp = svd(X[,order[1:10]])

nfolds number of training sets

rank array with elements indicating the number of confounding covariates to be used in the random effect.

nthreads number of threads to be used

# **Examples**

```
n = 300
p = 5000
X = matrix(sample(0:2, n*p, replace=TRUE), nrow=n)

dcmp = svd(X)

# simulate response
h_sq = .8
eta = dcmp$u[,1:2] %*% rgamma(2, 2, 1)
error_var = (1-h_sq) / h_sq * var(eta)
y = eta + rnorm(n, sd=sqrt(error_var))

# Get ordering based on marginal correlation
```

df.residual.lrgpr 7

```
i = order(cor(y, X)^2, decreasing=TRUE)
# Fit cross-validation
fit = cv.lrgpr( y ~ 1, features=X, order=i)
plot.cv.lrgpr(fit)
```

```
df.residual.lrgpr Residual Degrees-of-Freedom
```

#### **Description**

Residual df from fit of lrgpr

# Usage

```
df.residual.lrgpr(object, ...)
```

# Arguments

```
object model fit with lrgpr
... other arguments
```

error.bar

Plot Error Bars

# Description

Plot error bars for a confidence interval

#### Usage

```
error.bar(x, y, upper, lower = upper, length = 0.1, ...)
```

```
x x-axis position
y y-axis position
upper height of bar above y
lower height of bar below y
length horizontal length of the error bar
... arguments for arrows(...)
```

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getAlleleFreq

Calculate allele frequency

# Description

Calculate allele frequency

#### Usage

```
getAlleleFreq(X, nthreads = detectCores(logical = TRUE),
    progress = TRUE)
```

#### **Arguments**

X matrix where each column is a marker coded 0,1,2 or with dosage values in this

range

nthreads number of threads to use

progress show progress bar

getAlleleVariance EValuate variance for each column

## Description

EValuate variance for each column

#### Usage

```
getAlleleVariance(X,
  nthreads = detectCores(logical = TRUE),
  progress = TRUE)
```

# Arguments

X matrix where each column is a marker

nthreads number of threads to use

progress show progress bar

getMissingCount 9

| getMissingCount | Count missing values |
|-----------------|----------------------|
|                 |                      |

#### **Description**

Count missing values

#### Usage

```
getMissingCount(X,
  nthreads = detectCores(logical = TRUE),
  progress = TRUE)
```

# Arguments

X matrix where each column is a marker

nthreads number of threads to use progress show progress bar

glmApply

Fit standard linear or logistic model for many markers

# Description

'glmApply' is analogous to 'lrgprApply', but fits standard linear or logistic models for many markers

# Usage

```
glmApply(formula, features, terms = NULL,
  family = gaussian(), useMean = TRUE,
  nthreads = detectCores(logical = TRUE),
  univariateTest = TRUE, multivariateTest = FALSE,
  verbose = FALSE, progress = TRUE)
```

| formula standard linear modeling syntax as used in 'lm'. SNP is a place heach successive column of features | standard linear modeling syntax as used in 'lm'. SNP is a place holder for the each successive column of features |
|---|---|
|   | a matrix where the statistical model is evaluated with SNP if formula replace by each column successively         |
| terms   | indices of the coefficients to be tested. The indices corresponding to SNP are used if terms is not specified     |

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gaussian() for a continuous response, and binomial() to fit a logit model for a binary response

useMean if TRUE, replace missing entries with column mean. Otherwise, do not evaluate the model for that column

nthreads number of to use for parallel execution

univariateTest
 perform univariate hypothesis test for each response for each feature in the loop variable

multivariateTest
 perform multivariate hypothesis test for each response (if more than one) for

perform multivariate hypothesis test for each response (if more than one) for each feature. Note that the runtime is cubic in the number of response variables

verbose print additional information progress show progress bar

#### **Examples**

```
# Generate data
n = 100
p = 500
X = matrix(sample(0:2, n*p, replace=TRUE), nrow=n)
y = rnorm(n)
sex = as.factor(sample(1:2, n, replace=TRUE))
# Fit model for all markers
pValues = qlmApply(y \sim sex + sex:SNP, features=X, terms=c(3,4))
# Multivariate model
n = 100
p = 1000
m = 10
Y = matrix(rnorm(n*m), nrow=n, ncol=m)
X = matrix(rnorm(n*p), nrow=n, ncol=p)
res = glmApply( Y ~ SNP, features = X, terms=2, multivariateTest=TRUE)
# p-values for univariate hypothesis test of each feature against
 each response
res$pValues
# p-values for multivariate hypothesis test of each feature against
 all responses are the same time
# returns the results of the Hotelling and Pillai tests
res$pValues_mv
# The multivariate test for X[,1]
res$pValues_mv[1,]
# The result is the same as the standard tests in R
```

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```
fit = manova( Y ~ X[,1])
summary(fit, test="Hotelling-Lawley")
summary(fit, test="Pillai")
```

influence.lrgpr

Regression Diagnostics

# Description

Basic quantities for regression diagnostics from fit of lrgpr

#### Usage

```
influence.lrgpr(model, ...)
```

# Arguments

```
model model fit with lrgpr
... other arguments
```

leverage.lrgpr

Regression Diagnostics

# Description

Basic quantities for regression diagnostics from fit of lrgpr

# Usage

```
leverage.lrgpr(object)
```

```
object model fit with lrgpr
```

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```
lm.influence.lrgpr Regression Diagnostics
```

# Description

Basic quantities for regression diagnostics from fit of lrgpr

# Usage

```
lm.influence.lrgpr(object, ...)
```

# Arguments

```
object model fit with lrgpr
... other arguments
```

logLik.lrgpr

Extract Log-Likelihood

# Description

Log-Likelihood for model fit by lrgpr

# Usage

```
logLik.lrgpr(object, ...)
```

```
object model fit with lrgpr
... other arguments
```

loss.lrgpr

|--|

# Description

Compare observed and fitted response under some loss function

# Usage

```
loss.lrgpr(y, yhat, family)
```

#### **Arguments**

| У      | observed response   |
|--------|---|
| yhat   | fitted response   |
| family | "gaussian" or "binomial"  |
|        |   |
|        |   |
| lrgpr  | Fit a Low Rank Gaussian Process Regression (LRGPR)/Linear Mixed |
|        | Model (LMM)   |

# Description

'lrgpr' is used to fit LRGPR/LMM models that account for covariance in response values, but where the scale of the covariance is unknown. Standard linear modeling syntax is used for the model specification in addition to a covariance matrix or its eigen-decomposition.

# Usage

```
lrgpr(formula, decomp,
  rank = max(length(decomp$d), length(decomp$values)),
  delta = NULL, nthreads = detectCores(logical = TRUE),
  W_til = NULL, scale = TRUE)
```

| formula  | standard linear modeling syntax as used in 'lm'  |
|----------|--|
| decomp   | eigen-decomposition produced from eigen(K), where K is the covariance matrix. Or singular value decomposition $svd(X[,1:100])$ based on a subset of markers          |
| rank     | decomposition is truncated to the first rank eigen-vectors   |
| delta    | ratio of variance components governing the fit of the model. This should be estimated from a previous evaluation of 'lm' on the same response and eigendecomposition |
| nthreads | number of threads to use for parallel execution  |

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W\_til markers used to construct decomp that should now be removed from costruction of decomp. This is the proximal contamination term of Listgarten, et al. (2012) should W\_til be scaled and centered

#### Value

coefficients regression coefficients for each covariate

p.values p-values from Wald test of each coefficient

sd standard deviation of each coefficient estimate

sigSq\_e variance component

 $\sigma_e^2$ 

corresponding to the residual error

sigSq\_a variance component

 $\sigma_a^2$ 

corresponding the scale of the covariance, K

delta ratio of variance components:

 $\sigma_e^2/\sigma_a^2$ 

rank the rank of the random effect logLik log-likelihood of the model fit

fitted.values

estimated response values: y\_hat

alpha BLUP of the random effect

Sigma variance-covariate matrix of estimate of beta
hii diagonals of the matrix H such that y\_hat = Hy

y responses x design matrix

df effective degrees of freedom: trace(H) based on Hoffman (2013)

residuals residuals of model fit: y - y\_hat
AIC Akaike information criterion
BIC Bayesian information criterion
GCV generalized cross-validation
eigenVectors eigen-vectors in decomp
eigenValues eigen-values in decomp

df.residual n-ncol(X)

rank rank of decomposition used, where only non-negative eigen/singular values are

considered

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#### **Details**

'lrgpr' fits the model:

$$y = X\beta + \alpha + \epsilon$$

$$\alpha \sim N(0, K\sigma_a^2)$$

$$\epsilon \sim N(0, \sigma_e^2)$$

where

$$\delta = \sigma_e^2/\sigma_a^2$$

In practice the eigen-decomposition of K, and not K itself is required. The rank can be set to use only eigen-vectors 1:rank in the model.

This package allows hypothesis tests of single coefficients using fit\$p.values which fits a Wald test. Composite hypothesis tests of multiple coefficients are performed with wald(fit, terms=1:3).

Note that likelihood ratio tests with linear mixed models do not perform well and the resulting p-values often do not follow a uniform distribution under the null (Pinheiro and Bates, 2000). We strongly advise against using it with this model.

'lrgpr' uses the algorithm of Lippert, et al. (2011).

See Hoffman (2013) for an interpretation of the linear mixed model.

# References

Kang, H. M., et al. (2010) Variance component model to account for sample structure in genome-wide association studies. \_Nature Genetics\_ 42, 348-54

Lippert, C., et al. (2011) FaST linear mixed models for genome-wide association studies. \_Nature Methods\_ 9, 525-26

Listgarten, J., et al. (2012) Improved linear mixed models for genome-wide association studies. \_Nature Methods\_ 8, 833-5

Rasmussen, C. E. and Williams, C. K. I. (2006) Gaussian processes for machine learning. MIT Press

Pinheiro, J. C. and Bates, D. M. (2000) Mixed-Effects Models in S and S-PLUS. Springer, New York

Hoffman, G. E. (2013) Correcting for Population Structure and Kinship Using the Linear Mixed Model: Theory and Extensions. \_PLoS ONE\_ 8(10):e75707

Note that degrees freedom and some diagnostic statistics are not currently calculated with  $W_{\text{til}}$  is specified.

#### See Also

'wald', 'lrgprApply'

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#### **Examples**

```
# Generate random data
set.seed(1)
n < -200
y <- rnorm(n)
K <- crossprod( matrix(rnorm(n*1000), ncol=n) )</pre>
age <- rpois(n, 50)
sex <- as.factor(sample(1:2, n, replace=TRUE))</pre>
decomp <- eigen(K)
# Fit the model
fit <- lrgpr( y ~ sex + age, decomp)
# Print results
fit
# Print more detailed results
summary(fit)
# P-values for each covariate
fit$p.values
# Visualize fit of the model like for 'lm'
par(mfrow=c(2,2))
plot(fit)
# Composite hypothesis test using Wald's test
# Joint test of coefficients 2:3
wald( fit, terms=2:3)
```

lrgprApply

Fit a Low Rank Gaussian Process Regression (LRGPR)/Linear Mixed Model (LMM) for many markers

#### **Description**

'lrgprApply' is used to fit LRGPR/LMM models that account for covariance in response values, but where the scale of the covariance is unknown. It returns p-values equivalent to the results of lrgpr() and wald(), but is designed to analyze thousands of markers in a single function call.

#### Usage

```
lrgprApply(formula, features, decomp, terms = NULL,
  rank = max(length(decomp$d), length(decomp$values)),
  map = NULL, distance = NULL, dcmp_features = NULL,
  W_til = NULL, scale = TRUE, delta = NULL,
  reEstimateDelta = FALSE,
  nthreads = detectCores(logical = TRUE),
  verbose = FALSE, progress = TRUE)
```

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### **Arguments**

| standard linear modeling syntax as used in 'lm'. SNP is a place holder each successive column of features |   |
|---|---|
| features  | a matrix where the statistical model is evaluated with SNP if formula replace by each column successively   |
| decomp eigen-decomposition produced from eigen(K), where K is the covar                                   | eigen-decomposition produced from eigen(K), where K is the covariance matrix. Or singular value decomposition $svd(features[,1:100])$ based on a subset of markers  |
| terms   | indices of the coefficients to be tested. The indices corresponding to SNP are used if terms is not specified   |
| rank  | decomposition is truncated to the first rank eigen-vectors  |
| map   | p x 2 matrix where each entry corresponds to a marker in features. First column is the marker names, second columns is the genetic or physical location   |
| distance  | size of the proximal contamination window in units specifed by map.   |
| dcmp_feature  | S   |
|   | the indices in features of the markers used to construct dcmp   |
|   | and mandes in reactives of the manners used to construct demp   |
| W_til   | markers used to construct decomp that should now be removed from costruction of decomp. This is the proximal contamination term of Listgarten, et al. (2012)  |
| W_til   | markers used to construct decomp that should now be removed from costruction  |
| _   | markers used to construct decomp that should now be removed from costruction of decomp. This is the proximal contamination term of Listgarten, et al. (2012)  |
| scale   | markers used to construct decomp that should now be removed from costruction of decomp. This is the proximal contamination term of Listgarten, et al. (2012) should W_til be scaled and centered ratio of variance components governing the fit of the model. This should be estimated from a previous evaluation of 'lm' on the same response and eigendecomposition   |
| scale<br>delta  | markers used to construct decomp that should now be removed from costruction of decomp. This is the proximal contamination term of Listgarten, et al. (2012) should W_til be scaled and centered ratio of variance components governing the fit of the model. This should be estimated from a previous evaluation of 'lm' on the same response and eigendecomposition   |
| scale<br>delta  | markers used to construct decomp that should now be removed from costruction of decomp. This is the proximal contamination term of Listgarten, et al. (2012) should W_til be scaled and centered ratio of variance components governing the fit of the model. This should be estimated from a previous evaluation of 'lm' on the same response and eigendecomposition  1ta should delta be re-estimated for every marker. Note: reEstimateDelta=TRUE is             |
| scale delta reEstimateDe  | markers used to construct decomp that should now be removed from costruction of decomp. This is the proximal contamination term of Listgarten, et al. (2012) should W_til be scaled and centered ratio of variance components governing the fit of the model. This should be estimated from a previous evaluation of 'lm' on the same response and eigendecomposition  lta should delta be re-estimated for every marker. Note: reEstimateDelta=TRUE is much slower |

# **Examples**

```
# Generate data
n = 100
p = 500
X = matrix(sample(0:2, n*p, replace=TRUE), nrow=n)
y = rnorm(n)
sex = as.factor(sample(1:2, n, replace=TRUE))

K = tcrossprod(matrix(rnorm(n*n*3), nrow=n))
decomp = eigen(K, symmetric=TRUE)

# Fit null model
fit = lrgpr( y ~ sex, decomp)

# Fit model for all markers
pValues = lrgprApply( y ~ sex + sex:SNP, features=X, decomp, terms=c(3,4), delta=fit$delta)
```

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```
plot.criterion.lrgpr
```

Plot AIC/BIC/GCV values for lrgpr() model as rank changes

### Description

'plot.criterion.lrgpr' plots the criteria returned by 'criterion.lrgpr'

#### Usage

```
plot.criterion.lrgpr(x, col = rainbow(3), ...)
```

#### **Arguments**

```
x list returned by 'criterion.lrgpr'col array of 3 colors... other arguments
```

#### See Also

criterion.lrgpr

```
plot.cv.lrgpr
```

Plot Results of Cross-validation

#### **Description**

Plot results of 'cv.lrgpr', which fits cross-validation for multiple ranks of the LRGPR

#### Usage

```
plot.cv.lrgpr(x,
   ylim = c(min(x$cve - x$cvse), max(x$cve + x$cvse)),
   xlim = range(x$rank), pch = 20, col = "red",
   main = "Cross validation", xlab = "# of markers used",
   ylab = "Cross validation error", ...)
```

```
x result of cv.lrgpr
ylim limits of y-axis
xlim limits of x-axis
pch pch
col col
```

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```
main main

xlab xlab

ylab ylab

... other parameters fed to plot()
```

plot.lrgpr

Plot Diagnostics for an lrgpr Object

#### **Description**

Six plots (selectable by \"which\") are currently available: a plot of residuals against fitted values, a Scale-Location plot of sqrt(| residuals |) against fitted values, a Normal Q-Q plot, a plot of Cook's distances versus row labels, a plot of residuals against leverages, and a plot of Cook's distances against leverage/(1-leverage). By default, the first three and \"5\" are provided.

#### Usage

```
plot.lrgpr(x, which = c(1L:3L, 5L),
   caption = list("Residuals vs Fitted", "Normal Q-Q", "Scale-Location", "Cook's of panel = if (add.smooth) panel.smooth else points,
   sub.caption = NULL, main = "",
   ask = prod(par("mfcol")) < length(which) && dev.interactive(),
   ..., id.n = 3, labels.id = names(residuals(x)),
   cex.id = 0.75, qqline = TRUE, cook.levels = c(0.5, 1),
   add.smooth = getOption("add.smooth"),
   label.pos = c(4, 2), cex.caption = 1)</pre>
```

| Х           | lrgpr object.   |
|-------------|---|
| which       | if a subset of the plots is required, specify a subset of the numbers \"1:6\".  |
| caption     | captions to appear above the plots; \"character\" vector or \"list\" of valid graphics annotations, see \"as.graphicsAnnot\". Can be set to \"""\" or \"NA\" to suppress all captions.          |
| panel       | panel function. The useful alternative to \"points\", \"panel.smooth\" can be chosen by \"add.smooth = $TRUE$ \".   |
| sub.caption | common title-above the figures if there are more than one; used as \"sub\" (s.\"title\") otherwise. If \"NULL\", as by default, a possible abbreviated version of \"deparse(x\$call)\" is used. |
| main        | title to each plot-in addition to \"caption\".  |
| ask         | logical; if \"TRUE\", the user is _ask_ed before each plot, see \"par(ask=.)\".   |
|             | other parameters to be passed through to plotting functions.  |
| id.n        | number of points to be labelled in each plot, starting with the most extreme.   |

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| labels.id   | vector of labels, from which the labels for extreme points will be chosen. \"NULL\" uses observation numbers. |
|-------------|---|
| cex.id      | magnification of point labels.  |
| qqline      | logical indicating if a \"qqline()\" should be added to the normal Q-Q plot.                                  |
| cook.levels | levels of Cook's distance at which to draw contours.  |
| add.smooth  | logical indicating if a smoother should be added to most plots; see also \"panel\" above.                     |
| label.pos   | positioning of labels, for the left half and right half of the graph respectively, for plots 1-3.             |
| cex.caption | controls the size of \"caption\".   |

# See Also

plot.lm

| Predict response |
|------------------|
|------------------|

# Description

Predict response values after training with lrgpr. Leaving  $X_{test}$  and  $K_{test}$  as NULL returns the fitted values on the training set

# Usage

```
predict.lrgpr(object, X_test = NULL, K_test = NULL, ...)
```

| object | model fit from lrgpr on training samples                           |
|--------|--|
| X_test | design matrix of covariates for test samples                       |
| K_test | covariance matrix between samples in the test set and training set |
|        | other arguments  |

print.lrgpr 21

```
print.lrgpr
```

Print Values

# Description

Print details for fit from lrgpr

# Usage

```
print.lrgpr(x, ...)
```

# Arguments

```
x model fit from lrgpr... other arguments
```

# Description

Print summary for fit from lrgpr

# Usage

```
print.summary.lrgpr(x, ...)
```

# Arguments

```
x model fit from lrgpr
```

... other arguments

QQ\_plot

#### **Description**

QQ plot and lambda\_GC optimizd for large datasets.

# Usage

```
QQ_plot(p_values,
  col = rainbow(min(length(p_values), ncol(p_values))),
  main = "", pch = 20, errors = TRUE, lambda = TRUE,
  p_thresh = 1e-06, showNames = FALSE, ylim = NULL,
  xlim = NULL, plot = TRUE, new = TRUE,
  box.lty = par("lty"), collapse = FALSE, ...)
```

# Arguments

| p_values  | vector, matrix or list of p-values   |
|-----------|--|
| col       | colors corresponding to the number of columns in matrix, or entries in the list  |
| main      | title  |
| pch       | pch  |
| errors    | show 95% confidence interval   |
| lambda    | calculate and show genomic control lambda. Lambda_GC is calculated using the 'median' method on p-values $>$ p_thresh. |
| p_thresh  | Lambda_GC is calcualted using the 'median' method on p-values > p_thresh.  |
| showNames | show column names or list keys in the legend   |
| ylim      | ylim   |
| xlim      | xlim   |
| plot      | make a plot. If FALSE, returns lamda_GC values without making plot   |
| new       | make a new plot. If FALSE, overlays QQ over current plot   |
| box.lty   | box line type  |
| collapse  | combine entries in matrix or list into a single vector   |
|           | other arguments  |

#### **Examples**

```
p = runif(1e6)
QQ_plot(p)

# get lambda_GC values without making plot
lambda = QQ_plot(p, plot=FALSE)
```

read.fam 23

read.fam

Read plink FAM/TFAM files

#### **Description**

Read FAM/TFAM file into a dataframe. This function is the same as read.tfam

#### Usage

```
read.fam(file)
```

#### **Arguments**

file

location of FAM/TFAM file

read.tfam

Read plink FAM/TFAM files

# Description

Read FAM/TFAM file into a dataframe. This function is the same as read.fam

#### Usage

```
read.tfam(file)
```

# Arguments

file

location of FAM/TFAM file

residuals.lrgpr

Extract Model Residuals

# Description

Residuals fitted with lrgpr

# Usage

```
residuals.lrgpr(object, type = "working", ...)
```

# Arguments

```
object model fit with lrgpr
```

type the type of residual, but there is only one option here

... other arguments

24 summary.lrgpr

```
rstandard.lrgpr
```

Regression Deletion Diagnostics

#### **Description**

Basic quantities for regression deletion diagnostics from fit of lrgpr

# Usage

```
rstandard.lrgpr(model, ...)
```

# Arguments

```
model model fit with lrgpr
... other arguments
```

```
set_missing_to_mean
```

Replace Missing Values with Mean

#### **Description**

For each column, replace NA values with the column mean

#### Usage

```
set_missing_to_mean(A)
```

#### **Arguments**

Α

matrix

```
summary.lrgpr
```

Summarizing LRGPR / Linear Mixed Model Fits

#### **Description**

Print summary for fit from lrgpr

## Usage

```
summary.lrgpr(object, ...)
```

```
object model fit from lrgpr
... other arguments
```

vcov.lrgpr 25

vcov.lrgpr

Calculate Variance-Covariance Matrix for a lrgpr Object

# Description

Returns the variance-covariance matrix of the main parameters of a fitted model object

# Usage

```
vcov.lrgpr(object, ...)
```

# **Arguments**

object model fit with lrgpr
... other arguments

wald

Composite hypothesis test of multiple coefficients

# Description

'wald' performs a multi-dimensional Wald test against H0: beta\_i...beta\_j = 0 using the estimated coefficients and their variance-covariance matrix

#### Usage

```
wald(fit, terms)
```

#### **Arguments**

fit result of fitting with 'lrgpr'

terms indices of the coefficients to be tested

#### **Details**

The Wald statistic is

$$\beta_h^T \Sigma_h^{-1} \beta_h \sim \chi_{|h|}^2$$

where

h

specifies the coefficients being tested and

h

is the number of entries

#### See Also

'lrgpr'

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